#2



OIPE

RAW SEQUENCE LISTING

.

DATE: 03/19/2002

PATENT APPLICATION: US/09/824,647

TIME: 14:49:08

Input Set : N:\Crf3\RULE60\09824647.raw
Output Set: N:\CRF3\03192002\1824647.raw

```
1 <110> APPLICANT: Serrero, Ginette
      2 <120> TITLE OF INVENTION: 88 KDA TUMORIGENIC GROWTH FACTOR AND ANTAGONISTS
      3 <130> FILE REFERENCE: Z9996.488/P001-A
      4 <140> CURRENT APPLICATION NUMBER: 09/824,647
      5 <141> CURRENT FILING DATE: 2001-04-04
      7 <150> PRIOR APPLICATION NUMBER: EARLIER APPLICATION NUMBER: 08/991,862
    8 <151> PRIOR FILING DATE: EARLIER FILING DATE: 1998-08-17
     11 <150> PRIOR APPLICATION NUMBER: EARLIER APPLICATION NUMBER: 08/863,862
W--> 12 <151> PRIOR FILING DATE: EARLIER FILING DATE: 1997-05-23
     13 <160> NUMBER OF SEQ ID NOS: 17
     14 <170> SOFTWARE: PatentIn Ver. 2.0
                                                            ENTERED
     16 <210> SEQ ID NO: 1
     17 <211> LENGTH: 2137
     18 <212> TYPE: DNA
     19 <213> ORGANISM: Mouse epithelin/granulin
     20 <220> FEATURE:
     21 <221> NAME/KEY: CDS
     22 <222> LOCATION: (23)..(1789)
     23 <223> OTHER INFORMATION: The sequence is identical to that of the published
              mouse granulin except for one nucleotide (T
    25
              instead of G) at position 1071 of GP88 cDNA
              (position 1056 of mouse granulin).
     26
     27 <400> SEQUENCE: 1
     28
              eggacecega egeagacaga ee atg tgg gte etg atg age tgg etg gee tte.
                                       Met Trp Val Leu Met Ser Trp Leu Ala Phe
     29
     30
              gcg gca ggg ctg gta gcc gga aca cag tgt cca gat ggg cag ttc tgc
                                                                                 100
     31
              Ala Ala Gly Leu Val Ala Gly Thr Gln Cys Pro Asp Gly Gln Phe Cys
     32
     33
                               15
                                                   20
              cct gtt gcc tgc tgc ctt gac cag gga gga gcc aac tac agc tgc tgt
                                                                                 148
     35
              Pro Val Ala Cys Cys Leu Asp Gln Gly Gly Ala Asn Tyr Ser Cys Cys
     36
                           30
                                               35
                                                                                 196
     37
              aac cet ett etg gae aca tgg eet aga ata acg age eat eat eta gat
              Asn Pro Leu Leu Asp Thr Trp Pro Arg Ile Thr Ser His His Leu Asp
     38
     39
                                           50
                                                                                 244
              gge tee tge cag ace cat gge cac tgt cet get gge tat tet tgt ett
     40
              Gly Ser Cys Gln Thr His Gly His Cys Pro Ala Gly Tyr Ser Cys Leu
     41
     42
                   60
                                       65
     43
              ctc act gtg tct ggg act tcc agc tgc tgc ccg ttc tct aag ggt gtg
                                                                                 292
             Leu Thr Val Ser Gly Thr Ser Ser Cys Cys Pro Phe Ser Lys Gly Val
     44
     45
     46
              tot tgt ggt gat ggc tac cac tgc tgc ccc cag ggc ttc cac tgt agt
                                                                                 340
```

Ser Cys Gly Asp Gly Tyr His Cys Cys Pro Gln Gly Phe His Cys Ser

47

Input Set : N:\Crf3\RULE60\09824647.raw
Output Set: N:\CRF3\03192002\1824647.raw

48					95					100					105		
49	gca	gat	ggg	aaa	tcc	tgc	ttc	cag	atg	tca	gat	aac	ccc	ttg	ggt	gct	388
50						Cys											
51		_	_	110		_			115					120		•	
52	gtc	cag	tgt	cct	ggg	agc	cag	ttt	gaa	tgt	cct	gac	tct	gcc	acc	tgc	436
53	Val	Gln	Cys	Pro	Gly	Ser	Gln	Phe	Glu	Cys	Pro	Asp	Ser	Ala	Thr	Cys	
54			125					130					135				
55	tgc	att	atg	gtt	gat	ggt	tcg	tgg	gga	tgt	tgt	ccc	atg	ccc	cag	gcc	484
56	Cys	Ile	Met	Val	Asp	Gly	Ser	Trp	Gly	Cys	Cys	Pro	Met	Pro	Gln	Ala	
57		140					145					150					
58	tct	tgc	tgt	gaa	gac	aga	gtg	cat	tgc	tgt	CCC	cat	ggg	gcc	tcc	tgt	532
59	Ser	Cys	Cys	Glu	Asp	Arg	Val	His	Cys	Cys	Pro	His	Gly	Ala	Ser	Cys	
60	155					160					165					170	
61	gac	ctg	gtt	cac	aca	cga	tgc	gtt	tca	ccc	acg	ggc	acc	cac	acc	cta	580
62						Arg											
63					175					180					185		
64	cta	aag	aag	ttc	cct	gca	caa	aag	acc	aac	agc	gca	gtg	tct	ttg	cct	628
65						Āla											
66			_	190					195					200			
67	ttt	tct	gtc	gtg	tgc	cct	gat	gct	aag	acc	cag	tgt	ccc	gat	gat	tct	676
68						Pro											
69			205					210					215				
70	acc	tgc	tgt	gag	cta	ccc	act	ggg	aag	tat	ggc	tgc	tgt	cca	atg	ccc	724
71	Thr	Cys	Cys	Glu	Leu	Pro	Thr	Gly	Lys	Tyr	Gly	Cys	Cys	Pro	Met	Pro	
72		220	-				225	_				230					
73	aat	gcc	atc	tgc	tgt	tcc	gac	cac	ctg	cac	tgc	tgc	ccç	cag	gac	act	772
74	Asn	Ala	Ile	Cys	Cys	Ser	Asp	His	Leu	His	Cys	Cys	Pro	Gln	Asp	Thr	
75	235			_	_	240					245					250	
76	gta	tgt	gac	ctg	atc	cag	agt	aag	tgc	cta	tcc	aag	aac	tac	acc	acg	820
77	Val	Cys	Asp	Leu	Ile	Gln	Ser	Lys	Cys	Leu	Ser	Lys	Asn	Tyr	Thr	Thr	
78					255					260	•				265		
79	gat	ctc	ctg	acc	aag	ctg	cct	gga	tac	cca	gtg	aag	gag	gtg	aag	tgc	868
80	Asp	Leu	Leu	Thr	Lys	Leu	Pro	Gly	Tyr	Pro	Val	Lys	Glu	Val	Lys	Cys	
81				270					275					280			
82	gac	atg	gag	gtg	agc	tgc	cct	gaa	gga	tat	acc	tgc	tgc	cgc	ctc	aac	916
83	Asp	Met	Glu	Val	Ser	Cys	Pro	Glu	Gly	Tyr	Thr	Cys		Arg	Leu	Asn	
84			285					290					295				
85						tgc											964
86	Thr	Gly	Ala	\mathtt{Trp}	Gly	Cys	Cys	Pro	Phe	Ala	Lys	Ala	Val	Cys	Cys	Asp	
87		300					305					310					
88						tgc											1012
89	_	His	Ile	His	Cys	Cys	Pro	Ala	Gly	Phe		Cys	His	Thr	Glu	Lys	
90	315					320					325					330	
91						ggt											1060
92	Gly	Thr	Cys	Glu		Gly	Ile	Leu	Gln		Gly	\mathtt{Trp}	Met	Lys		Val	
93					335					340					345		
94						ctg											1108
95	Ile	Ala	Pro		Arg	Leu	Pro	Asp		Gln	Ile	Leu	Lys		Asp	Thr	
96				350					355					360			

Input Set : N:\Crf3\RULE60\09824647.raw
Output Set: N:\CRF3\03192002\I824647.raw

97 98	cct Pro																1156
99	110	C	365	пор			****9	370	110		11011		375	O _I B	0,0	270	
100	ctc	aat		aaa	gac	t.aa	ααc	•	. tat	ccc	atc	cca		act	gtc	tac	1204
101															Val		
102		380		0-1			385		. 012			390				-1-	
103	tac			aac	caq	cat			cct	caq	qqc	ttc	aca	tat	ctg	qct	1252
104	-		-		_		_	_		_				-	Leu		
105	395					400	-				405			_		410	
106	cag	qqq	tac	tgt	cag	aag	gga	gac	aca	atg	gtg	gct	ggc	ctg	gag	aag	1300
107															Glu		
108		_	_	-	415	_	_	_		420					425		
109	ata	cct	gcc	cgc	cag	aca	acc	ccg	cto	caa	att	gga	gat	atc	ggt	tgt	1348
110	Ile	Pro	Ala	Arg	Gln	Thr	Thr	Pro	Leu	Gln	Ile	Gly	Asp	Ile	Gly	Cys	
111				430					435	j				440			
112	gac	cag	cat	acc	agc	tgc	cca	gta	ggg	caa	acc	tgc	tgc	cca	agc	ctc	1396
113	Asp	Gln	His	Thr	Ser	Cys	Pro	Val	. Gly	Gln	Thr	Cys	Cys	Pro	Ser	Leu	
114			445					450)				455				
115	aag	gga	agt	tgg	gcc	tgc	tgc	cag	r ctg	ccc	cat	gct	gtg	tgc	tgt	gag	1444
116	Lys	Gly	Ser	\mathtt{Trp}	Ala	Cys	Cys	Glr	Leu	Pro	His	Ala	Val	Cys	Cys	Glu	
117		460					465					470					
118	_		_		_	_	_	_				_			aag		1492
119		Arg	Gln	His	Cys	Cys	Pro	Ala	Gly	y Tyr	Thr	Cys	Asn	Val	Lys		
120	475					480					485					490	
121			_		_	-	_	_							ctc		1540
122	Arg	Thr	Cys	Glu	_	Asp	Val	Asp	Phe			Pro	Pro	Val	Leu	Leu	
123					495					500					505		
124					_	-									cat		1588
125	Thr	Leu	GTA		Lys	Val	GLY	Asn			Cys	GLY	GIU		His	Pne	
126				510			+		515		~+	~~~	~~~	520		~~~	1626
127	_		_		-		_	_		_					tgg		1636
128 129	Cys	HIS	525		GIII	THE	Cys	530	_	ASP	ser	Ата	535		Trp	Ата	
130	taa	+ a+			ot a	220	aat			. +a+	202	a+			cac	tat	1684
131															His		1004
132	Cys	540		- y -	Lea	БyЗ	545		. Cyb	, cys	nig	550		**** 9	1115	CID	
133	tac			aac	ttc	cac			acc	. agg	σσа			tat	ttg	сда	1732
134	_						-								Leu		
135	555		017	011	1110	560	_	-		9	565		-1-	010		570	
136		aaσ	att	cct	cac			ato	r ttt	tta		gat	cca	atc	cca		1780
137															Pro		
138	-1-	-1-			575	-				580	_	•			585	_	
139	ccq	cta	ctg	taa	ggaa	qqq	ctac	agac	tt a	agga	actc	c ac	agto	ctgg			1829
140	-		Leu										-				
142					gggt	ac c	cact	acto	a gg	cctc	ccta	gcg	cctc	ctc	ccct	aacgtc	1889
143																actāaa	
144																ttcttc	
145																gagctt	
146	gct	tgtg	tgt	gtgt	gcgc	gt g	tgcg	tgtg	rt tg	rctcc	aata	aag	tttg	tac	gctt	tctgaa	2129

Input Set : N:\Crf3\RULE60\09824647.raw
Output Set: N:\CRF3\03192002\I824647.raw

147 149 <210>		aaaa ID 1		2													2137
		LENGTH: 589															
151 <212>	TYPE	TYPE: PRT															
152 <213>	> ORGANISM: Mouse epithelin/granulin																
153 <400>	SEQU	JENCI	Ξ: 2														
154	Met	\mathtt{Trp}	Val	Leu	Met	Ser	Trp	Leu	Ala		Ala	Ala	Gly	Leu		Ala	
155	1				5			_	_	10		_	_		15		
156	Gly	Thr	Gln		Pro	Asp	Gly	Gln		Cys	Pro	Val	Ala		Cys	Leu	
157	_			20		_	_		25	_	_	_	_	_ 30	_	_,	•
158	Asp	Gin	_	GTÄ	Āla	Asn	Tyr		Cys	Cys	Asn	Pro		Leu	Asp	Thr	
159	m	D	35	- 1-	m1	a	TT -	40	T	3	01	0	45		mh m	TT	
160	Trp		Arg	тте	Thr	ser		HIS	ьeu	Asp	GIY	ser 60	Cys	GIII	THE	HIS	
161	C1,,	50	Cvrc	Dro	Ala	C117	55	Sor	Cvc	Tou	T.OU		17 a 1	Sor	G1 v	mhr.	
162 163	65	птэ	Cys	PIO	АІС	70	тут	261	Cys	пеа	75	1111	Val	Ser	GIY	80	
164		Sor	Cve	Cve	Pro		Ser	T.vc	Glv	Va 1		Cvs	Glv	Δsn	Glv		
165	Der	DCI	Cys	Cys	85	riic	Der	цуз		90	001	Cys	O.J	пор	95	-7-	
166	His	Cvs	Cvs	Pro	Gln	Glv	Phe	His	Cvs		Ala	Asp	Glv	Lvs	_	Cvs	
167		012	0,0	100	0	0-1			105				1	110		-1-	
168	Phe	Gln	Met		Asp	Asn	Pro	Leu		Ala	Val	Gln	Cys	Pro	Gly	Ser	
169			115					120					125		_		
170	Gln	Phe	Glu	Cys	Pro	Asp	Ser	Ala	Thr	Cys	Cys	Ile	Met	Val	Asp	Gly	
171		130		-		_	135			_	_	140			_	_	
172	Ser	Trp	Gly	Cys	Cys	Pro	Met	Pro	Gln	Ala	Ser	Cys	Cys	Glu	Asp	Arg	
173	145					150					155					160	
174	Val	His	Cys	Cys	Pro	His	Gly	Ala	Ser	Cys	Asp	Leu	Val	His	Thr	Arg	
175				•	165					170					175		
176	Cys	Val	Ser		Thr	Gly	Thr	His		Leu	Leu	Lys	Lys		Pro	Ala	
177				180		_	_		185			_	· -	190	_		
178	Gln	Lys		Asn	Ser	Ala	Val		Leu	Pro	Phe	Ser		Val	Cys	Pro	
179	_		195	m1	a 1	_		200	•	a		G	205	01	T	D	
180	Asp		ьуs	Thr	Gln	Cys		Asp	Asp	ser	Thr	_	Cys	GIU	ьeu	Pro	
181	mbs	210	T *** G	П	Gly	Crra	215	Dro	Mot	Dro	N an	220	т1.	Cvc	Cvc	Sor	
182 183	225	СТУ	гуѕ	TAT	GIY	230	Cys	PIO	Met	PIO	235	нта	TIE	Cys	Cys	240	
184		Uic	T.011	Шie	Cys		Dro	Gln	Δen	Thr		Cve	Δen	T.e.11	Tle		
185	изр	птэ	пец	III	245	Cys	110	GIII	изр	250	V U I	Cys	пор	пси	255	0111	
186	Ser	Lvs	Cvs	Len	Ser	Lvs	Asn	Tvr	Thr		Asp	Leu	Leu	Thr		Leu	
187	DCI	_, _	0,0	260	501	2,0		-1-	265					270	-1-		
188	Pro	Glv	Tvr		Val	Lvs	Glu	Val		Cvs	Asp	Met	Glu	Val	Ser	Cys	
189		1	275			-1-		280					285			-	
190	Pro	Glu	Gly	Tyr	Thr	Cys	Cys		Leu	Asn	Thr	Gly	Ala	Trp	Gly	Cys	
191		290	-	•		-	295	•				300		_	-	_	
192	Cys		Phe	Ala	Lys	Ala	Val	Cys	Cys	Asp	Asp	His	Ile	His	Cys	Cys	
193	305				-	310		-	-		315					320	
194	Pro	Ala	${\tt Gly}$	Phe	Gln	Cys	His	Thr	Glu	Lys	Gly	Thr	Cys	Glu	Met	Gly	
195					325					330					335		
196	Ile	Leu	Gln	Val	Gly	Trp	Met	Lys	Lys	Val	Ile	Ala	Pro	Leu	Arg	Leu	

Input Set : N:\Crf3\RULE60\09824647.raw
Output Set: N:\CRF3\03192002\I824647.raw

```
340
                                           345
                                                                350
197
          Pro Asp Pro Gln Ile Leu Lys Ser Asp Thr Pro Cys Asp Asp Phe Thr
198
199
                                       360
                                                            365
200
          Arg Cys Pro Thr Asn Asn Thr Cys Cys Lys Leu Asn Ser Gly Asp Trp
201
              370
                                   375
          Gly Cys Cys Pro Ile Pro Glu Ala Val Cys Cys Ser Asp Asn Gln His
202
203
                              390
                                                   395
          Cys Cys Pro Gln Gly Phe Thr Cys Leu Ala Gln Gly Tyr Cys Gln Lys
204
                                               410
205
          Gly Asp Thr Met Val Ala Gly Leu Glu Lys Ile Pro Ala Arg Gln Thr
206
207
                                           425
208
          Thr Pro Leu Gln Ile Gly Asp Ile Gly Cys Asp Gln His Thr Ser Cys
209
                                       440
          Pro Val Gly Gln Thr Cys Cys Pro Ser Leu Lys Gly Ser Trp Ala Cys
210
211
                                   455
          Cys Gln Leu Pro His Ala Val Cys Cys Glu Asp Arg Gln His Cys Cys
212
213
                               470
                                                   475
          Pro Ala Gly Tyr Thr Cys Asn Val Lys Ala Arg Thr Cys Glu Lys Asp
214
215
                          485
                                               490
216
          Val Asp Phe Ile Gln Pro Pro Val Leu Leu Thr Leu Gly Pro Lys Val
                                           505
217
          Gly Asn Val Glu Cys Gly Glu Gly His Phe Cys His Asp Asn Gln Thr
218
                                                            525
219
220
          Cys Cys Lys Asp Ser Ala Gly Val Trp Ala Cys Cys Pro Tyr Leu Lys
221
                                   535
                                                        540
          Gly Val Cys Cys Arg Asp Gly Arg His Cys Cys Pro Gly Gly Phe His
222
223
                               550
                                                   555
          Cys Ser Ala Arg Gly Thr Lys Cys Leu Arg Lys Lys Ile Pro Arg Trp
224
                                               570
225
                          565
          Asp Met Phe Leu Arg Asp Pro Val Pro Arg Pro Leu Leu
226
227
                      580
                                           585
229 <210> SEQ ID NO: 3
230 <211> LENGTH: 19
231 <212> TYPE: PRT
232 <213> ORGANISM: mouse granulin
233 <220> FEATURE:
234 <221> NAME/KEY: PEPTIDE
235 <222> LOCATION: (1)..(19)
236 <223> OTHER INFORMATION: Internal peptide of mouse GP88 used to raise the
237
          antisera against the GP88 used in the
          immunoaffinity step.
239 <400> SEQUENCE: 3
          Lys Lys Val Ile Ala Pro Arg Arg Leu Pro Asp Pro Gln Ile Leu Lys
240
241
                                                10
242
          Ser Asp Thr
244 <210> SEQ ID NO: 4
245 <211> LENGTH: 12
246 <212> TYPE: PRT
247 <213> ORGANISM: mouse granulin
```

VERIFICATION SUMMARY
PATENT APPLICATION: US/09/824,647
DATE: 03/19/2002
TIME: 14:49:09

Input Set : N:\Crf3\RULE60\09824647.raw
Output Set: N:\CRF3\03192002\1824647.raw

L:8 M:256 W: Invalid Numeric Header Field, Wrong Prior FILING DATE:YYYY-MM-DD
L:12 M:256 W: Invalid Numeric Header Field, Wrong Prior FILING DATE:YYYY-MM-DD
L:304 M:257 W: Feature value mis-spelled or invalid, <221> Name/Key for SEQ ID#:8
L:316 M:257 W: Feature value mis-spelled or invalid, <221> Name/Key for SEQ ID#:9
L:328 M:257 W: Feature value mis-spelled or invalid, <221> Name/Key for SEQ ID#:10
L:339 M:257 W: Feature value mis-spelled or invalid, <221> Name/Key for SEQ ID#:11
L:350 M:257 W: Feature value mis-spelled or invalid, <221> Name/Key for SEQ ID#:12
L:361 M:257 W: Feature value mis-spelled or invalid, <221> Name/Key for SEQ ID#:13
L:372 M:257 W: Feature value mis-spelled or invalid, <221> Name/Key for SEQ ID#:14
L:383 M:257 W: Feature value mis-spelled or invalid, <221> Name/Key for SEQ ID#:15